

A computational framework for particle and whole cell tracking applied to a real biological dataset

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Objectives

The human fibrosarcoma cell line HT-1080 obtained from DSMZ, Germany

F. Yang, C. Venkataraman, V. Styles, V. Kuttenger, E. Horn, Z. von Guttenger, A. Madzvamuse, Journal of Biomechanics,

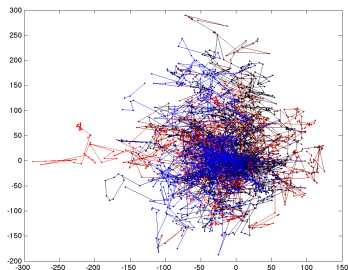
Accepted for publication, <http://dx.doi.org/10.1016/j.jbiomech.2016.02.008>, 2016.

- Identification from phase contrast microscopy
- Particle tracking
- Whole cell tracking for morphological changes
 - Optimal control of phase field formulations of geometric evolution laws
 - Efficient solver
 - Applications

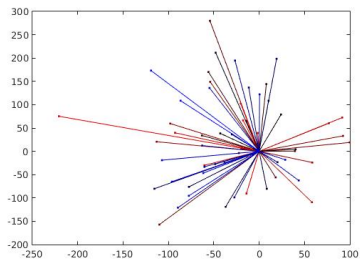
Techniques based upon background removal

Directions of migration

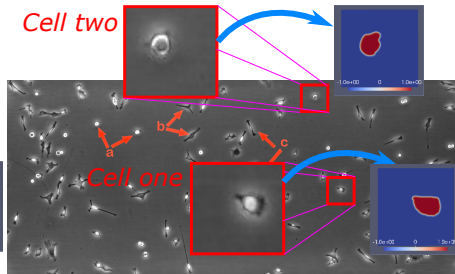
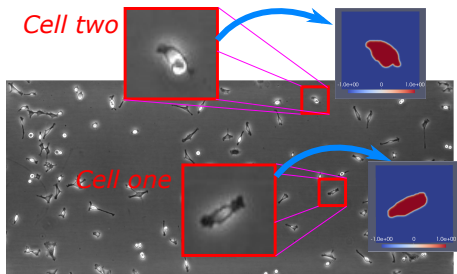
Spider plot



Star plot



Individual cells



Our optimal control model

The mass constrained mean curvature flow with forcing:

$$\begin{cases} \mathbf{V}(\mathbf{x}, t) &= (-\sigma H(\mathbf{x}, t) + \eta(\mathbf{x}, t) + \lambda_V(t)) \mathbf{v}(\mathbf{x}, t) \text{ on } \Gamma(t), t \in (0, T], \\ \Gamma(0) &= \Gamma^0. \end{cases}$$

The phase-field approximation of the above equation - Allen-Cahn:

$$\begin{cases} \partial_t \phi(\mathbf{x}, t) &= \Delta \phi(\mathbf{x}, t) - \frac{1}{\epsilon^2} G'(\phi(\mathbf{x}, t)) - \frac{1}{\epsilon} (\eta(\mathbf{x}, t) - \lambda(t)) \text{ in } \Omega \times (0, T], \\ \nabla \phi \cdot \boldsymbol{\nu}_\Omega &= 0 \text{ on } \partial\Omega \times (0, T], \\ \phi(\cdot, 0) &= \phi^0 \text{ in } \Omega. \end{cases}$$

Our optimal control model cont.

The objective functional:

$$J(\phi, \eta) = \frac{1}{2} \int_{\Omega} (\phi(\mathbf{x}, T) - \phi_{obs}(\mathbf{x}))^2 d\mathbf{x} + \frac{\theta}{2} \int_0^T \int_{\Omega} \eta(\mathbf{x}, t)^2 d\mathbf{x} dt,$$

and now we solve the minimisation problem:

$$\min_{\eta} J(\phi, \eta), \text{ with } J \text{ given above.}$$

Our optimal control model cont.

The adjoint equation to help computing the derivative of the objective functional:

$$\begin{cases} \partial_t p(\mathbf{x}, t) = -\Delta p(\mathbf{x}, t) + \epsilon^{-2} G''(\phi(\mathbf{x}, t)) p(\mathbf{x}, t) & \text{in } \Omega \times [0, T), \\ p(\mathbf{x}, T) = \phi(\mathbf{x}, T) - \phi_{obs}(\mathbf{x}) & \text{in } \Omega, \end{cases}$$

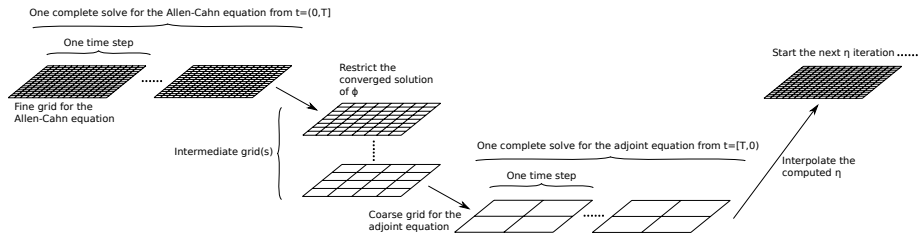
and we update the control as

$$\eta^{\ell+1} = \eta^\ell - \alpha \left(\theta \eta^\ell + \frac{1}{\epsilon} p^\ell \right).$$

Numerical challenges

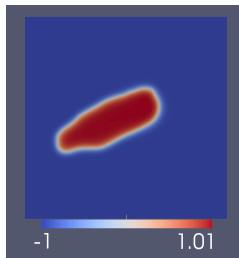
- Number of time steps
- Memory requirement (let's consider double precision and 100 time steps)
 - 2-D: 512^2 requires 0.4 gigabytes
 - 3-D: 512^3 requires 215 gigabytes

Two-grid solution strategy

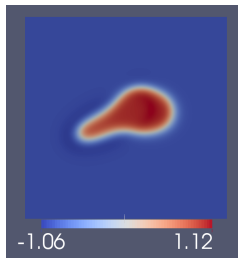


Cell one

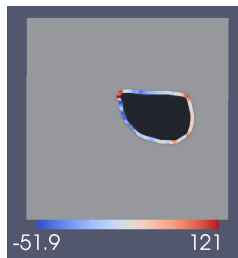
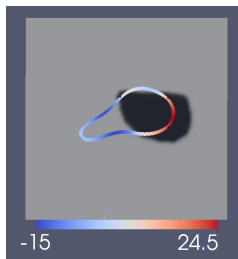
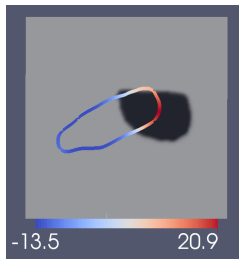
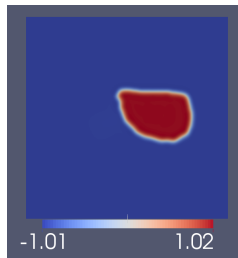
$t=0$



$t=T/2$



$t=T$

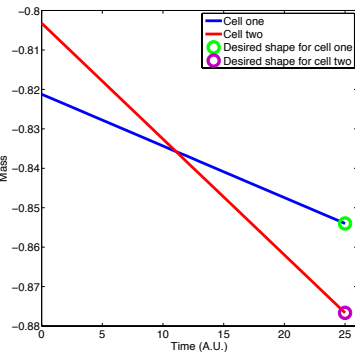


Cell one video

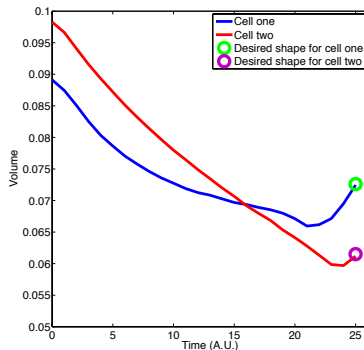
Cell two video

Analysis through tracking morphological changes

$$\int_{\Omega} \phi dx$$



$$\int_{\{\phi > 0\}} 1 dx$$



Real world example (2)

Euler number for topological changes

We compute this Euler number for these time steps with an "optimized" control η :

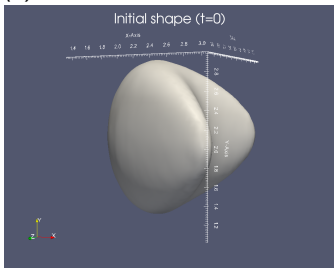
$$\mathcal{X} = \frac{1}{2\pi(a-b)} \int_{\Omega(a,b)} \left(-\Delta\phi + \frac{\nabla|\nabla\phi|^2 \cdot \nabla\phi}{2|\nabla\phi|^2} \right) dx.$$

Q. Du et al. *J. Appl. Math.*, 2005

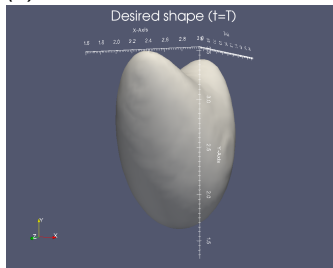
Real world example (2)

A 3-D example

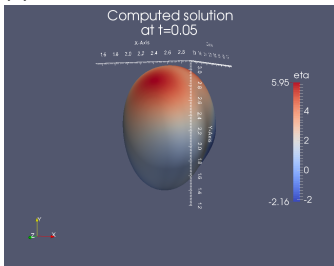
(a)



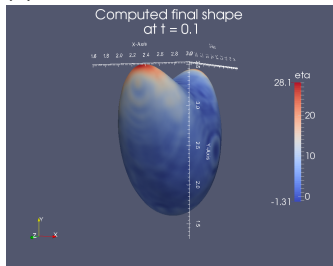
(b)



(c)



(d)



A 3-D example video

F.W. Yang, C.E. Goodyer, M.E. Hubbard and P.K. Jimack

“An Optimally Efficient Technique for the Solution of Systems of Nonlinear Parabolic Partial Differential Equations”

AiES in review, 2015

F. Yang, C. Venkataraman, V. Styles and A. Madzvamuse

“A Robust and Efficient Adaptive Multigrid Solver for the Optimal Control of Phase Field Formulations of Geometric Evolution Laws”

CiCP in review, 2015

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“A Computational Framework for Particle and Whole Cell Tracking Applied to a Real Biological Dataset”

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